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## TRANSMITTAL FORM

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Total Number of Pages in This Submission

Application Number	09/699,003
Filing Date	October 26, 2000
First Named Inventor	M. Rigdon Lentz
Art Unit	3762
Examiner Name	Patricia Bianco
Attorney Docket Number	LEN 101 CIP CON

### ENCLOSURES (Check all that apply)

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Remarks

~~05/03/2005 HDESTA1 00000001 503129 09699003~~  
~~01 EC:1806 180.00 DA~~

### SIGNATURE OF APPLICANT, ATTORNEY, OR AGENT

Firm Name	Pabst Patent Group LLP		
Signature			
Printed name	Tiffany B. Salmon		
Date	April 29, 2005	Reg. No.	55,589

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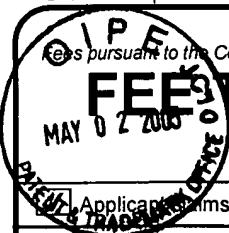
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 <p>Effective on 12/08/2004. Fees pursuant to the Consolidated Appropriations Act, 2005 (H.R. 4818).</p> <h1 style="text-align: center;">FEE TRANSMITTAL</h1> <h2 style="text-align: center;">For FY 2005</h2>		<p style="text-align: center;"><b>Complete if Known</b></p> <table border="1"> <tr><td>Application Number</td><td>09/699,003</td></tr> <tr><td>Filing Date</td><td>October 26, 2000</td></tr> <tr><td>First Named Inventor</td><td>M. Rigdon Lentz</td></tr> <tr><td>Examiner Name</td><td>Patricia Bianco</td></tr> <tr><td>Art Unit</td><td>3762</td></tr> <tr><td>Attorney Docket No.</td><td>LEN 101 CIP CON</td></tr> </table>		Application Number	09/699,003	Filing Date	October 26, 2000	First Named Inventor	M. Rigdon Lentz	Examiner Name	Patricia Bianco	Art Unit	3762	Attorney Docket No.	LEN 101 CIP CON
Application Number	09/699,003														
Filing Date	October 26, 2000														
First Named Inventor	M. Rigdon Lentz														
Examiner Name	Patricia Bianco														
Art Unit	3762														
Attorney Docket No.	LEN 101 CIP CON														
<input checked="" type="checkbox"/> Application claims small entity status. See 37 CFR 1.27															
TOTAL AMOUNT OF PAYMENT	(\$ 180.00)														

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**FEE CALCULATION****1. BASIC FILING, SEARCH, AND EXAMINATION FEES**

Application Type	FILING FEES		SEARCH FEES		EXAMINATION FEES		Fees Paid (\$)
	Fee (\$)	Small Entity Fee (\$)	Fee (\$)	Small Entity Fee (\$)	Fee (\$)	Small Entity Fee (\$)	
Utility	300	150	500	250	200	100	
Design	200	100	100	50	130	65	
Plant	200	100	300	150	160	80	
Reissue	300	150	500	250	600	300	
Provisional	200	100	0	0	0	0	

**2. EXCESS CLAIM FEES**

Fee Description	Fee (\$)	Small Entity Fee (\$)
Each claim over 20 or, for Reissues, each claim over 20 and more than in the original patent	50	25
Each independent claim over 3 or, for Reissues, each independent claim more than in the original patent	200	100
Multiple dependent claims	360	180

<u>Total Claims</u>	<u>Extra Claims</u>	<u>Fee (\$)</u>	<u>Fee Paid (\$)</u>	<u>Multiple Dependent Claims</u>	
<u>15</u> - 20 or HP =	<u>0</u>	x <u>0.00</u>	= <u>0.00</u>	<u>Fee (\$)</u>	<u>Fee Paid (\$)</u>
HP = highest number of total claims paid for, if greater than 20					
<u>Indep. Claims</u>	<u>Extra Claims</u>	<u>Fee (\$)</u>	<u>Fee Paid (\$)</u>		
<u>2</u> - 3 or HP =	<u>0</u>	x <u>0.00</u>	= <u>0.00</u>		
HP = highest number of independent claims paid for, if greater than 3					

**3. APPLICATION SIZE FEE**

If the specification and drawings exceed 100 sheets of paper, the application size fee due is \$250 (\$125 for small entity) for each additional 50 sheets or fraction thereof. See 35 U.S.C. 41(a)(1)(G) and 37 CFR 1.16(s).

Total Sheets	Extra Sheets	Number of each additional 50 or fraction thereof	Fee (\$)	Fee Paid (\$)
_____ - 100 = _____	/ 50 = _____	(round up to a whole number) x _____	= _____	

**4. OTHER FEE(S)**


Non-English Specification, \$130 fee (no small entity discount)

Other: Information Disclosure Statement

Fees Paid (\$)

\$180.00

**SUBMITTED BY**

Signature		Registration No. (Attorney/Agent)	55,589	Telephone	(404) 879-2153
Name (Print/Type)	Tiffany B. Salmon	Date	April 29, 2005		

This collection of information is required by 37 CFR 1.136. The information is required to obtain or retain a benefit by the public which is to file (and by the USPTO to process) an application. Confidentiality is governed by 35 U.S.C. 122 and 37 CFR 1.14. This collection is estimated to take 30 minutes to complete, including gathering, preparing, and submitting the completed application form to the USPTO. Time will vary depending upon the individual case. Any comments on the amount of time you require to complete this form and/or suggestions for reducing this burden, should be sent to the Chief Information Officer, U.S. Patent and Trademark Office, U.S. Department of Commerce, P.O. Box 1450, Alexandria, VA 22313-1450. DO NOT SEND FEES OR COMPLETED FORMS TO THIS ADDRESS. SEND TO: Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313-1450.

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IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Applicants: M. Rigdon Lentz

Serial No.: 09/699,003

Art Unit: 3762

Filed: October 26, 2000

Examiner: P. Bianco

For: *METHOD AND COMPOSITIONS FOR TREATMENT OF CANCERS*

Commissioner for Patents  
P.O. Box 1450  
Alexandria, VA 22313-1450

**SUPPLEMENTAL INFORMATION DISCLOSURE STATEMENT**

Sir:

Pursuant to the duty of disclosure under 37 C.F.R. §1.56 and 37 C.F.R. §1.97, Applicants submit a Supplemental Information Disclosure Statement, including fifteen (15) pages of Form PTO-1449, and copies of the one hundred and ninety-four (194) documents cited therein.

This Information Disclosure Statement is being filed under 37 C.F.R. § 1.97(C), after a first Office Action on the merits. The Commissioner is authorized to charge \$180.00, the fee set forth under 37 CFR § 1.17(p), to Account No. 50-3129. It is believed that no additional fee is required with this submission. However, should an additional fee be required, the Commissioner is hereby authorized to charge any fees to Deposit Account No. 50-3129.

05/03/2005 HDESTA1 00000001 09699003

01 FC:1806 180.00 DA

### U.S. Patent Applications

<u>Number</u>	<u>Publication Date</u>	<u>Inventor</u>	<u>Class/Subclass</u>
2001/0010818	08-02-2001	Engle, et al.	
2001/0039392	11-08-2001	Strahilevitz	
2002/0019603	02-14-2002	Strahilevitz	
2002/0058031	05-16-2002	Tung, et al.	
2002/0086276	07-04-2002	Srivastava	
2002/0107469	08-08-2002	Bolan, et al.	
2002/0111577	08-15-2002	Sirimanne, et al.	
2002/0119147	08-29-2002	Howell, et al.	
2002/0159995	10-31-2002	Brady, et al.	
2002/0183677	12-05-2002	Chang, et al.	
2002/0187069	12-12-2002	Levin, et al.	
2002/0197249	12-26-2002	Brady, et al.	
2002/0197250	12-26-2002	Brady, et al.	
2002/0197251	12-26-2002	Brady, et al.	
2003/0073822	04-17-2003	Lofling, et al.	
2003/0118584	06-26-2003	Glenn, et al.	
2003/0125657	07-03-2003	Koll, et al.	
2003/0127390	07-10-2003	Davis, Jr.	
2003/0133929	07-17-2003	Cham	
2003/0138349	07-24-2003	Robinson, et al.	
2003/0148404	08-07-2003	Michaelson	
2003/0163077	08-28-2003	Kim, et al.	
2003/0195452	10-16-2003	Hunley, et al.	
2003/0215443	11-20-2003	Coffey, et al.	
2004/0044301	03-04-2004	Levin, et al.	
2004/0054315	03-18-2004	Levin, et al.	

### U.S. Patents

<u>Number</u>	<u>Issue Date</u>	<u>Patentee</u>	<u>Class/Subclass</u>
4,116,589	09-26-1978	Rishton	
4,189,470	02-19-1980	Rose	
4,191,182	03-04-1980	Popovich, et al.	
4,350,156	09-21-1982	Malchesky, et al.	
4,362,155	12-07-1982	Skurkovich	604/6.04
4,375,414	03-01-1983	Strahilevitz	
4,581,010	04-08-1986	Skurkovich, et al.	
4,605,394	08-12-1986	Skurkovich	
4,620,977	11-04-1986	Strahilevitz	

U.S.S.N.: 09/699,003  
 Filed: October 26, 2000  
 SUPPLEMENTAL INFORMATION  
 DISCLOSURE STATEMENT

4,633,417	12-30-1986	Wilburn, et al.
4,634,417	01-06-1987	Korec
4,664,913	05-12-1987	Mielke, et al.
4,801,449	01-31-1989	Balint, Jr., et al.
4,813,924	03-21-1989	Strahilevitz
4,824,432	04-25-1989	Skurkovich, et al.
4,834,973	05-30-1989	Strahilevitz
4,865,841	09-12-1989	Balint, Jr., et al.
4,963,265	10-16-1990	Okarma, et al.
5,037,645	08-06-1991	Strahilevitz
5,037,649	08-06-1991	Balint, Jr., et al.
5,078,673	01-07-1992	Abrams
5,356,374	10-18-1994	Hogan, et al.
5,605,690	02-25-1997	Jacobs, et al.
5,626,843	05-06-1997	Skurkovich, et al.
5,643,732	07-01-1997	Strahilevitz
5,705,615	01-06-1998	Lim, et al.
5,730,713	03-24-1998	Okarma, et al.
5,753,227	05-19-1998	Strahilevitz
5,840,588	11-24-1998	Strahilevitz
5,869,047	02-09-1999	Blake
5,888,511	03-30-1999	Skurkovich, et al.
5,910,252	06-08-1999	Truitt, et al.
6,039,946	03-21-2000	Strahilevitz
6,197,289	03-06-2001	Wirt, et al.
6,221,614	04-24-2001	Prusiner, et al.
6,245,038	06-12-2001	Borberg, et al.
6,264,623	07-24-2001	Strahilevitz
6,287,516	09-11-2001	Matson, et al.
6,428,790	08-06-2002	Boyd
6,432,405	08-13-2002	Weinberg, et al.
6,528,057	03-04-2003	Ambrus, et al.
6,561,997	05-13-2003	Weitzel, et al.
6,569,112	05-27-2003	Strahilevitz
6,602,502	08-05-2003	Strahilevitz
6,602,993	08-05-2003	Wallach, et al.
6,607,501	08-19-2003	Gorsuch
6,607,723	08-19-2003	Good, et al.
6,627,151	09-30-2003	Borberg, et al.
6,630,315	10-07-2003	Miwa, et al.
6,676,622	01-13-2004	Strahilevitz
6,685,664	02-03-2004	Levin, et al.

RE 31,688	09-25-1984	Popovich, et al.
RE 36,755	06-27-2000	Smith, et al.

### Foreign Documents

<u>Number</u>	<u>Publication Date</u>	<u>Patentee</u>	<u>Country</u>
3302384	01-25-1983	Lysaght, et al.	DE
0076665	01-14-1987	E.I. Du Pont De Nemours and Company	EP
2136314	09-19-1984	Gelman Sciences, Inc.	GB
56092824	07-27-1981	Terumo Corporation	JP
WO 79/01121	12-27-1979	The Government of the United States of America	PCT

### Publications

ABBRUZZESE, et al., "A phase II trial of recombinant human interferon-gamma and recombinant tumor necrosis factor in patients with advanced gastrointestinal malignancies: Results of a trial terminated by excessive toxicity" *Journal of Biological Response Modifiers* 9:522-527 (1990).

ADOLF AND APFLER, "A monoclonal antibody-based enzyme immunoassay for quantitation of human tumor necrosis factor binding protein I, a soluble fragment of the 60kDa TNF receptor, in biological fluids" *J. Immunol. Meth.* 143:127-136 (1991).

AJANI, et al., "Phase I and II studies of the combination of recombinant human interferon- $\gamma$  and 5-fluorouracil in patients with advanced colorectal carcinoma" *Journal of Biological Response Modifiers* 8:140-146 (1989).

ALBERTINI, et al., "Limiting dilution analysis of lymphokine-activated killer cell precursor frequencies in peripheral blood lymphocytes of cancer patients receiving interleukin-2 therapy" *Journal of Biological Response Modifiers* 9:456-462 (1990).

AREND, "Inhibiting the effects of cytokines in human diseases" *Adv. Int. Med.* 40:365-394 (1995).

AVNER, et al., "Therapeutic murine monoclonal antibodies developed for individual cancer patients" *Journal of Biological Response Modifiers* 8:25-36 (1989).

BALCEWICZ-SABLINSKA, et al., "Pathogenic *Mycobacterium tuberculosis* evades apoptosis of host macrophages by release of TNF-R2, resulting in inactivation of TNF- $\alpha$ " *J. Immunol.* 161:2636-2641 (1998).

BALIKO, et al., "Th2 biased immune response in cases with active *Mycobacterium tuberculosis* infection and tuberculin anergy" *FEMS Immunol. Med. Micro.* 22:188-204 (1998).

BERMUDEZ AND CHAMPSI, "Infection with *Mycobacterium avium* induces production of interleukin-10 (IL-10), and administration of anti-IL-10 antibody is associated with enhanced resistance to infection in mice" *Infect. Immun.* 61:3093-3097 (1993).

BEUTLER AND CERAMI, "The biology of cachectin/TNF-A primary mediator of the host response" *Ann. Rev. Immunol.* 7:625-655 (1989).

BLAUER, et al., "Modulation of the antilisterial activity of human blood-derived macrophages by activating and deactivating cytokines" *J. Interferon Cytokine Res.* 15:105-114 (1995).

BOMAN, et al., "Phase I study of recombinant gamma-interferon (rIFN- $\gamma$ )" *Journal of Biological Response Modifiers* 7:438-446 (1988).

BRUNTSCH, et al., "phase II study of recombinant human interferon-  $\gamma$  in metastatic renal cell carcinoma" *Journal of Biological Response Modifiers* 9:335-338 (1990).

BUKOWSKI, et al., "Phase I trial of continuous infusion of recombinant interleukin-2 and intermittent recombinant interferon- $\alpha_{2a}$ . Clinical effects" *Journal of Biological Response Modifiers* 9:538-545 (1990).

CAULFIELD, et al., "Phase II-Ib trial of an anti- $G_{D3}$  monoclonal antibody in combination with interferon- $\alpha$  in patients with malignant melanoma" *Journal of Biological Response Modifiers* 9:319-328 (1990).

CHAMBRIER, et al., "Hormonal and metabolic effects of chronic interleukin-2 infusion in cancer patients" *Journal of Biological Response Modifiers* 9:251-255 (1990).

CHOUAIB, et al., "More insights into the complex physiology of TNF" *Immunol. Today* 12(5):141-145 (1991).

COCLET-NININ, et al., "Interferon-beta not only inhibits interleukin-1 $\beta$  and tumor necrosis factor- $\alpha$  but stimulates interleukin-1 receptor antagonist production in human peripheral blood mononuclear cells" *Eur. Cytokine Network* 8(4):345-349 (1997).

COLMAN, et al., Hemostasis and Thrombosis: Basic Principles and Clinical Practice 2nd. Edition (Colman, et al., eds.) pp. 63-67 J.B. Lippincott: Philadelphia, PA, 1987.

COX, et al., "Phase II study of human lymphoblastoid interferon in patients with multiple myeloma" *Journal of Biological Response Modifiers* 7:318-325 (1988).

CREAVEN, et al., "Initial clinical trial of the macrophage activator muramyl tripeptide-phosphatidylethanolamine encapsulated in liposomes in patients with advanced cancer" *Journal of Biological Response Modifiers* 9:492-498 (1990).

CROGHAN, et al., "A phase I trial of recombinant interferon- $\alpha$  and  $\alpha$ -difluoromethylornithine in metastatic melanoma" *Journal of Biological Response Modifiers* 7:409-415 (1988).

D'ANDREA, et al., "Interleukin 10 (IL-10) inhibits human lymphocyte interferon  $\gamma$  production by suppressing natural killer cell stimulatory factor/IL-12 synthesis in accessory cells" *J. Exp. Med.* 178:1041-1048 (1993).

DIMERY, et al., "Recombinant interferon-  $\gamma$  in the treatment of recurrent nasopharyngeal carcinoma" *Journal of Biological Response Modifiers* 8:221-226 (1989).

DINARELLO, "Induction of interleukin-I and interleukin-I receptor antagonist" *Sem. In Oncol.* 24(no. 3, Suppl. 9):81-93 (1997).

DUPERE, et al., "Patterns of cytokines released by peripheral blood leukocytes of normal donors and cancer patients during interleukin-2 activation *in vitro*" *Journal of Biological Response Modifiers* 9:140-148 (1990).

ENGELHARDT, et al, "Biological response to intravenously administered endotoxin in patients with advanced cancer" *Journal of Biological Response Modifiers* 9:480-491 (1990)

ENGELMANN, et al., "Two tumor necrosis factor-binding proteins purified from human urine" *J. Biol. Chem.* 265(3):1531-1536 (1990).

ERIKS AND EMERSON, "Temporal effect of tumor necrosis factor alpha on murine macrophages infected with *Mycobacterium avium*" *Infect. Immun.* 65(6):2100-2106 (1997).

ETGES AND MULLER, "Progressive disease or protective immunity to *Leishmania* major infection: the result of a network of stimulatory and inhibitory interactions" *J. Mol. Med.* 76:372-390 (1998).

FAREED, et al., "Novel antigenic markers of human tumor regression" *Journal of Biological Response Modifiers* 7:11-23 (1988).

FAVROT, et al., "Functional and immunophenotypic modifications induced by interleukin-2 did not predict response to therapy in patients with renal cell carcinoma" *Journal of Biological Response Modifiers* 9:167-177 (1990).

FERNANDES AND BALDWIN, "Interleukin-10 downregulates protective immunity to *Brucella abortus*" *Infect. Immun.* 63:1130-1133 (1995).



FOON, et al., "A prospective randomized trial of  $\alpha_2$ B-interferon/ $\gamma$ -interferon or the combination in advanced metastatic renal cell carcinoma" *Journal of Biological Response Modifiers* 7:540-545 (1988).

FROST, et al., "Interleukin-6 induction by a muramyltripeptide derivative in cancer patients" *Journal of Biological Response Modifiers* 9:160-166 (1990).

GADDUCCI, et al., "Serum levels of soluble receptors for tumor necrosis factor (p55 and p75 sTNFr) in endometrial cancer" *Anticancer Res.* 16:3125-3128 (1996).

GILL, et al., "Interferon-alpha maintenance therapy after cytotoxic chemotherapy for treatment of acquired immunodeficiency syndrome-related kaposi's sarcoma" *Journal of Biological Response Modifiers* 9:512-516 (1990).

GREENBLATT, et al., "The type B receptor for tumor necrosis factor-alpha" *Blood* 80:1339-1346 (1992).

GUSTAVSON, et al., "Pharmacokinetics of teceleukin (Recombinant human interleukin-2) after intravenous or subcutaneous administration to patients with cancer" *Journal of Biological Response Modifiers* 8:440-449 (1989).

HANDZEL, et al., "Immunomodulation of T cell deficiency in humans by thymic humoral factor: from crude extract to synthetic thymic humoral factor- $\gamma$ 2" *Journal of Biological Response Modifiers* 9:269-278 (1990).

HANK, et al., "Depressed *in vitro* T cell responses concomitant with augmented interleukin-2 responses by lymphocytes from cancer patients following *in vivo* treatment with interleukin-2" *Journal of Biological Response Modifiers* 9:5-14 (1990).

HERCEND, et al., "Immunotherapy with lymphokine-activated natural killer cells and recombinant interleukin-2: A feasibility trial in metastatic renal cell carcinoma" *Journal of Biological Response Modifiers* 9:546-555 (1990).

HERRMANN, et al., "Stimulation of granulopoiesis in patients with malignancy by recombinant human granulocyte-macrophage colony-stimulating factor: Assessment of two routes of administration" *Journal of Biological Response Modifiers* 9:475-479 (1990).

HERTLER, et al., "A phase I study of T101-ricin A chain immunotoxin in refractory chronic lymphocytic leukemia" *Journal of Biological Response Modifiers* 7:97-113 (1988).

HIMMLER, et al., "Molecular cloning and expression of human and rat tumor necrosis factor receptor chain (p60) and its soluble derivative, tumor necrosis factor-binding protein" *DNA and Cell Biol.* 9(10):705-715 (1990).

JACOBS, et al., "minimal antigenicity of intron A in human recipients demonstrated by three analytical methods" *Journal of Biological Response Modifiers* 7:447-456 (1988).

JAKOBSEN, et al., "Decreased antitoxic activities among children with clinical episodes of malaria" *Infect. Immun.* 66(4):1654-1659 (1998).

JAKSCHIES, et al., "Emergence and decay of the human Ms homolog in cancer patients during and after interferon- $\alpha$  therapy" *Journal of Biological Response Modifiers* 9:305-312 (1990).

KALMANTI, et al., "Serum levels of tumor necrosis factor soluble interleukin 2 receptor as markers of disease activity and prognosis in childhood leukemia and lymphoma" *Int. J. Hematol.* 57:147-152 (1993).

KAUFMANN, et al., "T cells and cytokines in intracellular bacterial infections: experiences with *Mycobacterium bovis* BCG" *Ciba Fdn. Symp.* 195:123-132 (1995).

KELLOKUMPU-LEHTINEN, et al., "Recombinant interferon- $\alpha$ 2a and vinblastine in advanced renal cell cancer: A clinical phase I-II study" *Journal of Biological Response Modifiers* 9:439-444 (1990).

KESSLER, "Adsorptive plasma treatment: Optimization of extracorporeal devices and systems" *Blood Purification* 11:150-157 (1993).

KHAZAELI, et al., "Initial evaluation of a human immunoglobulin M monoclonal antibody (HA-1A) in humans" *Journal of Biological Response Modifiers* 9:178-184 (1990).

KOLITZ, et al., "Phase I trial of recombinant interleukin-2 and cyclophosphamide: Augmentation of cellular immunity and T-cell mitogenic response with long term administration of rIL-2" *Journal of Biological Response Modifiers* 7:457-472 (1988).

KRIGEL, et al., "Treatment of epidemic kaposi's sarcoma with a combination of interferon-alpha 2b and etoposide" *Journal of Biological Response Modifiers* 7:359-364 (1988).

LANTZ, et al., "Infusion of tumor necrosis factor (TNF) causes an increase in circulating TNF-binding protein in humans" *Cytokine* 2(6):402-406 (1990).

LASZLO, et al., "Phase I studies of recombinant interferon- $\gamma$ " *Journal of Biological Response Modifiers* 9:185-193 (1990).

LAUCELLA, et al., "Papel de las citoquinas en la resistencia y patologia durante la infeccion con *Trypanosoma cruzi*" *Revista Argentina de Microbiologia*, 28:99-109 (1996).

LENTZ, et al., "Apheresis of low molecular weight protein fraction and the onset of labor" *Journal of Clinical Apheresis* 5:62-67 (1990).

LENTZ, "The phylogeny of oncology" *Mol. Biother.* 2:137-144 (1990).

LETTERIO AND ROBERTS, "Regulation of immune responses by TGF- $\beta$ " *Ann. Rev. Immuno.* 16:137-161 (1998).

LITTON, et al., "Biological and clinical effects of the oral immunomodulator 3,6'-Bis(2-piperidinoethoxy)acridine trihydrochloride in patients with malignancy" *Journal of Biological Response Modifiers* 9:61-70 (1990).

LUCEY, et al., "Type 1 and type 2 cytokine dysregulation in human infectious, neoplastic, and inflammatory diseases" *Clin. Micro. Rev.* 9(4):532-562 (1996).

MAAS, et al., "Interleukin-2 in cancer treatment: disappointing or (still) promising? A review" *Cancer Immunol. Immunother.* 36:141-148 (1993).

MACA, "Inhibition of the growth of lewis lung carcinoma by indomethacin in conventional, nude, and beige mice" *Journal of Biological Response Modifiers* 9:568-580 (1990).

MARSHALL, et al., "Effects of coumarin (12-benzopyrone) and cimetidine on peripheral blood lymphocytes, natural killer cells, and monocytes in patients with advanced malignancies" *Journal of Biological Response Modifiers* 8:62-69 (1989).

MARSHALL, et al., "Treatment of renal cell carcinoma with daily low-dose alpha-interferon" *Journal of Biological Response Modifiers* 8:453-461 (1989).

MILES, et al., "induction of soluble tumour necrosis factor receptors during treatment with interleukin-2" *Brit. J. Cancer* 66:1195-1199 (1992).

MITTELMAN, et al., "Treatment of patients with advanced cancer using multiple long-term cultured lymphokine-activated killer (LAK) cell infusions and recombinant human interleukin-2" *Journal of Biological Response Modifiers* 8:468-478 (1989).

MUSIANI, et al., "Effect of low doses of interleukin-2 injected perilymphatically and peritumorally in patients with advanced primary head and neck squamous cell carcinoma" *Journal of Biological Response Modifiers* 8:571-578 (1989).

NEIDHART, "Phase I study of recombinant methionyl human consensus interferon (r-metHuIFN-Con)" *Journal of Biological Response Modifiers* 7:240-248 (1988).

ORATZ, et al. "Induction of tumor-infiltrating lymphocytes in human malignant melanoma metastases by immunization to melanoma antigen vaccine" *Journal of Biological Response Modifiers* 8:355-358 (1989).

ORATZ, et al., "Antimelanoma monoclonal antibody-ricin A chain immunoconjugate (XMMME-001-RTA) plus cyclophosphamide in the treatment of metastatic malignant melanoma: Results of a phase II trial" *Journal of Biological Response Modifiers* 9:345-354 (1990).

PAIS, et al., "Pharmacokinetics of recombinant interleukin-2 in children with malignancies: A pediatric oncology group study" *Journal of Biological Response Modifiers* 9:517-521 (1990).

PAOLOZZI, et al., "Phase I trial of recombinant interleukin-2 and recombinant  $\beta$ -interferon in refractory neoplastic diseases" *Journal of Biological Response Modifiers* 8:122-139 (1989).

PEREZ, et al., "A phase I trial of recombinant human gamma interferon (IFN- $\gamma_{4A}$ ) in patients with advanced malignancy" *Journal of Biological Response Modifiers* 7:309-317 (1988).

QUESADA, et al., "Recombinant interferon alpha and gamma in combination as treatment for metastatic renal cell carcinoma" *Journal of Biological Response Modifiers* 7:234-239 (1988).

RABINOWITZ, et al., "Hemolytic anemia in a cancer patient treated with recombinant interferon- $\gamma$ " *Journal of Biological Response Modifiers* 9:256-259 (1990).

REIMANN, et al., "Suppression of the immune response by micro-organisms" *Scand. J. Immunol.* 31:543-546 (1990).

RIFFKIN, et al., "Defense against the immune barrage: helminth survival strategies" *Immunol. Cell Bio* 74:564-574 (1996)

ROMANI, et al., "T helper cell dichotomy to *Candida albicans*: implications for pathology" *Immunol. Res.* 14:148-162 (1995).

ROSENTHAL, et al., "The *in vitro* function of lymphocytes from 25 cancer patients receiving four to seven consecutive days of recombinant IL-2" *Journal of Biological Response Modifiers* 7:123-139 (1988).

RYBAK, et al., "Interferon therapy of relapsed and refractory hodgkin's disease: Cancer and leukemia group B study 8652" *Journal of Biological Response Modifiers* 9:1-4 (1990).

SARNA, et al., "Systemic administration of recombinant methionyl human interleukin-2 (Ala 125) to cancer patients: Clinical results" *Journal of Biological Response Modifiers* 8:16-24 (1989).

SARNA, et al., "A pilot study of intralymphatic interleukin-2. II. Clinical and biological effects" *Journal of Biological Response Modifiers* 9:81-86 (1990).

SARTHOU, et al., "Prognostic value of anti-*Plasmodium falciparum*-specific immunoglobulin G3, cytokines, and their soluble receptors in west African patients with severe malaria" *Infect. Immun.* 65(8):3271-3276 (1997).

SATO, et al., "induction of bone formation in an adenoid cystic carcinoma of the maxillary sinus by adoptive immunotherapy involving intra-arterial injection of lympholine-activated killer cells and recombinant interleukin-2 in combination with radiotherapy" *Journal of Biological Response Modifiers* 9:329-334 (1990).

SCHAADT, et al., "Phase II study of recombinant human tumor necrosis factor in colorectal carcinoma" *Journal of Biological Response Modifiers* 9:247-250 (1990).

SCHALL, et al., "molecular cloning and expression of a receptor for human tumor necrosis factor" *Cell* 61:361-370 (1990).

SCHEITHAUER, et al., "Combined  $\alpha$ -2C-interferon/VMCP polychemotherapy versus VMCP polychemotherapy a induction therapy in multiple myeloma: A prospective randomized trial" *Journal of Biological Response Modifiers* 8:109-115 (1989).

SCHILLER, et al., "A phase I trial of interferon- $\alpha$ -2a plus cyclophosphamide, vincristine, prednisone, and doxorubicin" *Journal of Biological Response Modifiers* 8:252-261 (1989).

SECKINGER, et al., "Purification and biologic characterization of a specific tumor necrosis factor  $\alpha$  inhibitor" *J. Biol. Chem.* 264(20):11966-11973 (1989).

SEIGLER, et al., "Melanoma patient antibody responses to melanoma tumor-associated antigens defined by murine monoclonal antibodies" *Journal of Biological Response Modifiers* 8:37-52 (1989).

SHAU, et al., "A pilot study of intralymphatic interleukin-2. I. Cytotoxic and surface marker changes of peripheral blood lymphocytes" *Journal of Biological Response Modifiers* 9:71-80 (1990).

SIDHU AND BOLLON, "Tumor necrosis factor activities and cancer therapy – A perspective" *Pharmacol. Ther.* 57:79-128 (1993).

SELING, et al., "Immunosuppressive roles for IL-10 and IL-4 in human infection" *J. Immunol.* 150(12):5501-5510 (1993 ).

SPRIGGS, "One step ahead of the game: Viral immunomodulatory molecules" *Ann. Rev. Immunol.* 14:101-130 (1996).

STEGE, et al., "Long-term remission in a patient with erythroleukemia following interferon- $\alpha$  treatment" *Journal of Biological Response Modifiers* 8:351-354 (1989).

STEINMETZ, et al., "Phase I study of 24-hour continuous intravenous infusion of recombinant human tumor necrosis factor" *Journal of Biological Response Modifiers* 7:417-423 (1988).

SZOL, et al., "A phase I study of high-dose interleukin-2 in combination with interferon  $\alpha_2$ " *Journal of Biological Response Modifiers* 9:529-537 (1990).

TRIGG, et al., " $\alpha$ -interferon therapy for lymphoproliferative disorders developing in two children following bone marrow transplants" *Journal of Biological Response Modifiers* 8:603-613 (1989).

TRINCHIERI, et al., "Cytokine cross-talk between phagocytic cells and lymphocytes: Relevance for differentiation/Activation of phagocytic cells and regulation of adaptive immunity" *J. Cell. Biochem.* 53:301-308 (1993).

TRUMP et al., "Interferon- $\alpha$ -n1 and continuous infusion vinblastine for treatment of advanced renal cell carcinoma" *Journal of Biological Response Modifiers* 9:108-111 (1990).

UMIEL, et al., "Recombinant interleukin-2-activated intracavitary lymphocytes: Phenotypic characteristics and effector function" *Journal of Biological Response Modifiers* 8:409-421 (1989).

VON HOFF, et al., "Phase II evaluation of recombinant  $\gamma$ -interferon in patients with advanced pancreatic carcinoma: A southwest oncology group study" *Journal of Biological Response Modifiers* 9:584-587 (1990).

WALSH, et al., "Phase I study of the combination of alpha-2 interferon and cisplatin" *Journal of Biological Response Modifiers* 8:11-15 (1989).

WEIL-HILLMAN, et al., "Transient decrease in IL-2 responsive lymphocytes 24 hours after initiation of continuous IL-2 infusion in cancer patients" *Journal of Biological Response Modifiers* 7:424-437 (1988).

WHITEHEAD, et al., "A phase II trial of recombinant tumor necrosis factor in patients with metastatic colorectal adenocarcinoma: A southwest oncology group study" *Journal of Biological Response Modifiers* 9:588-591 (1990).

YELAVARTHI, et al., "Analysis of p60 and p80 tumor necrosis factor-alpha" *American Journal of Pathology* 143:1131-1141 (1993).

ZAMKOFF, et al., "A Phase I trial of subcutaneously administered recombinant tumor necrosis factor to patients with advanced malignancy" *Journal of Biological Response Modifiers* 8:539-552 (1989).

### Remarks

This statement should not be interpreted as a representation that an exhaustive search has been conducted or that no better art exists. Moreover, Applicants invite the Examiner to make an independent evaluation of the cited art to determine its relevance to the subject matter of the present application. Applicants are of the opinion that their claims patentably distinguish over the art referred to herein, either alone or in combination.

Respectfully submitted,




Tiffany B. Salmon  
Reg. No. 55,589

Dated: April 29, 2005

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		Filing Date	October 26, 2000
		First Named Inventor	M. Rigdon Lentz
		Group Art Unit	3762
		Examiner Name	P. Bianco
Sheet 1 of 15	Attorney Docket Number	LEN 101 CIP CON	

U.S. PATENT DOCUMENTS						
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		Number	Kind Code <sup>2</sup> (if known)			
		4,116,589		Rishton	09-26-1978	
		4,189,470		Rose	02-19-1980	
		4,191,182		Popovich, et al.	03-04-1980	
		4,350,156		Malchesky, et al.	09-21-1982	
		4,362,155		Skurkovich	12-07-1982	
		4,375,414		Strahilevitz	03-01-1983	
		4,581,010		Skurkovich, et al.	04-08-1986	
		4,605,394		Skurkovich	08-12-1986	
		4,620,977		Strahilevitz	11-04-1986	
		4,633,417		Wilburn, et al.	12-30-1986	
		4,634,417		Korec	01-06-1987	
		4,664,913		Mielke, et al.	05-12-1987	
		4,801,449		Balint, Jr., et al.	01-31-1989	
		4,813,924		Strahilevitz	03-21-1989	

FOREIGN PATENT DOCUMENTS								
Examiner Initials*	Cite No. <sup>1</sup>	Foreign Patent Document			Name of Patentee or Applicant of Cited Document	Date of Publication of Cited Document MM-DD-YYYY	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear	T <sup>6</sup>
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		DE	3302384		Lysaght, et al.	01-25-1983		
		EP	0076665		E.I. Du Pont De Nemours	01-14-1987		
		GB	2136314		Gelman Sciences, Inc.	09-19-1984		
		JP	56092824		Terumo Corporation	07-27-1981		
		PCT	WO 79/01121		The Government of the United States of America	12-27-1979		
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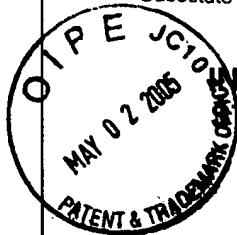






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		4,824,432		Skurkovich, et al.	04-25-1989	
		4,834,973		Strahilevitz	05-30-1989	
		4,865,841		Balint, Jr., et al.	09-12-1989	
		4,963,265		Okarma, et al.	10-16-1990	
		5,037,645		Strahilevitz	08-06-1991	
		5,037,649		Balint, Jr., et al.	08-06-1991	
		5,078,673		Abrams	01-07-1992	
		5,356,374		Hogan, et al.	10-18-1994	
		5,605,690		Jacobs, et al.	02-25-1997	
		5,626,843		Skurkovich, et al.	05-06-1997	
		5,643,732		Strahilevitz	07-01-1997	
		5,705,615		Lim, et al.	01-06-1998	
		5,730,713		Okarma, et al.	03-24-1998	
		5,753,227		Strahilevitz	05-19-1998	
		5,840,588		Strahilevitz	11-24-1998	
		5,869,047		Blake	02-09-1999	
		5,888,511		Skurkovich, et al.	03-30-1999	
		5,910,252		Truitt, et al.	06-08-1999	
		6,039,946		Strahilevitz	03-21-2000	
		6,197,289		Wirt, et al.	03-06-2001	
		6,221,614		Prusiner, et al.	04-24-2001	
		6,245,038		Borberg, et al.	06-12-2001	
		6,264,623		Strahilevitz	07-24-2001	
		6,287,516		Matson, et al.	09-11-2001	
		6,428,790		Boyd	08-06-2002	
		6,432,405		Weinberg, et al.	08-13-2002	
		6,528,057		Ambrus, et al.	03-04-2003	

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Date Considered

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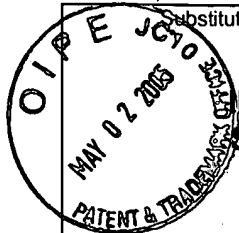
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## U.S. PATENT DOCUMENTS

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		Number	Kind Code <sup>2</sup> (if known)			
		6,561,997		Weitzel, et al.	05-13-2003	
		6,569,112		Strahilevitz	05-27-2003	
		6,602,502		Strahilevitz	08-05-2003	
		6,602,993		Wallach, et al.	08-05-2003	
		6,607,501		Gorsuch	08-19-2003	
		6,607,723		Good, et al.	08-19-2003	
		6,627,151		Borberg, et al.	09-30-2003	
		6,630,315		Miwa, et al.	10-07-2003	
		6,676,622		Strahilevitz	01-13-2004	
		6,685,664		Levin, et al.	02-03-2004	
		RE 31,688		Popovich, et al.	09-25-1984	
		RE 36,755		Smith, et al.	06-27-2000	
		2001/0010818		Engle, et al.	08-02-2001	
		2001/0039392		Strahilevitz	11-08-2001	
		2002/0019603		Strahilevitz	02-14-2002	
		2002/0058031		Tung, et al.	05-16-2002	
		2002/0086276		Srivastava	07-04-2002	
		2002/0107469		Bolan, et al.	08-08-2002	
		2002/0111577		Sirimanne, et al.	08-15-2002	
		2002/0119147		Howell, et al.	08-29-2002	
		2002/0159995		Brady, et al.	10-31-2002	
		2002/0183677		Chang, et al.	12-05-2002	
		2002/0187069		Levin, et al.	12-12-2002	
		2002/0197249		Brady, et al.	12-26-2002	
		2002/0197250		Brady, et al.	12-26-2002	
		2002/0197251		Brady, et al.	12-26-2002	
		2003/0073822		Lofling, et al.	04-17-2003	
		2003/0118584		Glenn, et al.	06-26-2003	

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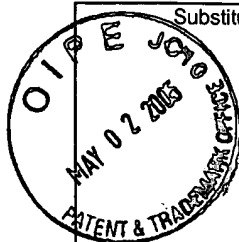
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		2003/0125657		Koll, et al.	07-03-2003	
		2003/0127390		Davis, Jr.	07-10-2003	
		2003/0133929		Cham	07-17-2003	
		2003/0138349		Robinson, et al.	07-24-2003	
		2003/0148404		Michaelson	08-07-2003	
		2003/0163077		Kim, et al.	08-28-2003	
		2003/0195452		Hunley, et al.	10-16-2003	
		2003/0215443		Coffey, et al.	11-20-2003	
		2004/0044301		Levin, et al.	03-04-2004	
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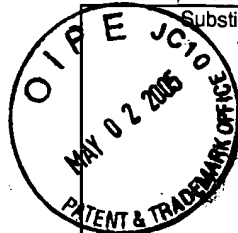
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		ABBRUZZESE, et al., "A phase II trial of recombinant human interferon-gamma and recombinant tumor necrosis factor in patients with advanced gastrointestinal malignancies: Results of a trial terminated by excessive toxicity" <i>Journal of Biological Response Modifiers</i> 9:522-527 (1990).	
		ADOLF AND APFLER, "A monoclonal antibody-based enzyme immunoassay for quantitation of human tumor necrosis factor binding protein I, a soluble fragment of the 60kDa TNF receptor, in biological fluids" <i>J. Immunol. Meth.</i> 143:127-136 (1991).	
		AJANI, et al., "Phase I and II studies of the combination of recombinant human interferon- $\gamma$ and 5-fluorouracil in patients with advanced colorectal carcinoma" <i>Journal of Biological Response Modifiers</i> 8:140-146 (1989).	
		ALBERTINI, et al., "Limiting dilution analysis of lymphokine-activated killer cell precursor frequencies in peripheral blood lymphocytes of cancer patients receiving interleukin-2 therapy" <i>Journal of Biological Response Modifiers</i> 9:456-462 (1990).	
		AREND, "Inhibiting the effects of cytokines in human diseases" <i>Adv. Int. Med.</i> 40:365-394 (1995).	
		AVNER, et al., "Therapeutic murine monoclonal antibodies developed for individual cancer patients" <i>Journal of Biological Response Modifiers</i> 8:25-36 (1989).	
		BALCEWICZ-SABLINSKA, et al., "Pathogenic <i>Mycobacterium tuberculosis</i> evades apoptosis of host macrophages by release of TNF-R2, resulting in inactivation of TNF- $\alpha$ " <i>J. Immunol.</i> 161:2636-2641 (1998).	
		BALIKO, et al., "Th2 biased immune response in cases with active <i>Mycobacterium tuberculosis</i> infection and tuberculin anergy" <i>FEMS Immunol. Med. Micro.</i> 22:188-204 (1998).	
		BERMUDEZ AND CHAMPSI, "Infection with <i>Mycobacterium avium</i> induces production of interleukin-10 (IL-10), and administration of anti-IL-10 antibody is associated with enhanced resistance to infection in mice" <i>Infect. Immun.</i> 61:3093-3097 (1993).	
		BEUTLER AND CERAMI, "The biology of cachectin/TNF-A primary mediator of the host response" <i>Ann. Rev. Immunol.</i> 7:625-655 (1989).	

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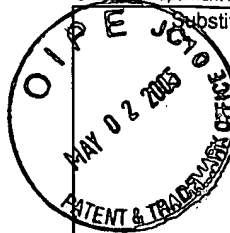
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## **Complete if Known**

Application Number	09/699,003
Filing Date	October 26, 2000
First Named Inventor	M. Rigdon Lentz
Group Art Unit	3762
Examiner Name	P. Bianco
Attorney Docket Number	LEN 101 CIP CON

Sheet 6 of 15

## **OTHER ART – NON PATENT LITERATURE DOCUMENTS**

Examiner's Initials*	Cite No. <sup>1</sup>	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published	T <sup>2</sup>
		BLAUER, et al., "Modulation of the antilisterial activity of human blood-derived macrophages by activating and deactivating cytokines" <i>J. Interferon Cytokine Res.</i> 15:105-114 (1995).	
		BOMAN, et al., "Phase I study of recombinant gamma-interferon (rIFN-γ)" <i>Journal of Biological Response Modifiers</i> 7:438-446 (1988).	
		BRUNTSCH, et al., "phase II study of recombinant human interferon- γ in metastatic renal cell carcinoma" <i>Journal of Biological Response Modifiers</i> 9:335-338 (1990).	
		BUKOWSKI, et al., "Phase I trial of continuous infusion of recombinant interleukin-2 and intermittent recombinant interferon-α <sub>2a</sub> . Clinical effects" <i>Journal of Biological Response Modifiers</i> 9:538-545 (1990).	
		CAULFIELD, et al., "Phase I1-Ib trial of an anti-G <sub>D3</sub> monoclonal antibody in combination with interferon-α in patients with malignant melanoma" <i>Journal of Biological Response Modifiers</i> 9:319-328 (1990).	
		CHAMBRIER, et al., "Hormonal and metabolic effects of chronic interleukin-2 infusion in cancer patients" <i>Journal of Biological Response Modifiers</i> 9:251-255 (1990).	
		CHOUAIB, et al., "More insights into the complex physiology of TNF" <i>Immunol. Today</i> 12(5):141-145 (1991).	
		COCLET-NININ, et al., "Interferon-beta not only inhibits interleukin-1β and tumor necrosis factor-α but stimulates interleukin-1 receptor antagonist production in human peripheral blood mononuclear cells" <i>Eur. Cytokine Network</i> 8(4):345-349 (1997).	
		COLMAN, et al., <i>Hemostasis and Thrombosis: Basic Principles and Clinical Practice</i> 2nd. Edition (Colman, et al., eds.) pp. 63-67 J.B. Lippincott: Philadelphia, PA, 1987.	
		COX, et al., "Phase II study of human lymphoblastoid interferon in patients with multiple myeloma" <i>Journal of Biological Response Modifiers</i> 7:318-325 (1988).	
		CREAVEN, et al., "Initial clinical trial of the macrophage activator muramyl tripeptide-phosphatidylethanolamine encapsulated in liposomes in patients with advanced cancer" <i>Journal of Biological Response Modifiers</i> 9:492-498 (1990).	

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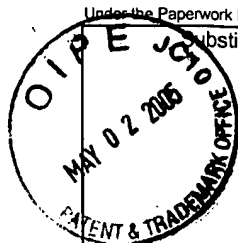
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		CROGHAN, et al., "A phase I trial of recombinant interferon- $\alpha$ and $\alpha$ -difluoromethylornithine in metastatic melanoma" <i>Journal of Biological Response Modifiers</i> 7:409-415 (1988).	
		D'ANDREA, et al., "Interleukin 10 (IL-10) inhibits human lymphocyte interferon $\gamma$ production by suppressing natural killer cell stimulatory factor/IL-12 synthesis in accessory cells" <i>J. Exp. Med.</i> 178:1041-1048 (1993).	
		DIMERY, et al., "Recombinant interferon- $\gamma$ in the treatment of recurrent nasopharyngeal carcinoma" <i>Journal of Biological Response Modifiers</i> 8:221-226 (1989).	
		DINARELLO, "Induction of interleukin-I and interleukin-I receptor antagonist" <i>Sem. In Oncol.</i> 24(no. 3, Suppl. 9):81-93 (1997).	
		DUPERE, et al., "Patterns of cytokines released by peripheral blood leukocytes of normal donors and cancer patients during interleukin-2 activation <i>in vitro</i> " <i>Journal of Biological Response Modifiers</i> 9:140-148 (1990).	
		ENGELHARDT, et al, "Biological response to intravenously administered endotoxin in patients with advanced cancer" <i>Journal of Biological Response Modifiers</i> 9:480-491 (1990)	
		ENGELMANN, et al., "Two tumor necrosis factor-binding proteins purified from human urine" <i>J. Biol. Chem.</i> 265(3):1531-1536 (1990).	
		ERIKS AND EMERSON, "Temporal effect of tumor necrosis factor alpha on murine macrophages infected with <i>Mycobacterium avium</i> " <i>Infect. Immun.</i> 65(6):2100-2106 (1997).	
		ETGES AND MULLER, "Progressive disease or protective immunity to <i>Leishmania</i> major infection: the result of a network of stimulatory and inhibitory interactions" <i>J. Mol. Med.</i> 76:372-390 (1998).	
		FAREED, et al., "Novel antigenic markers of human tumor regression" <i>Journal of Biological Response Modifiers</i> 7:11-23 (1988).	
		FAVROT, et al., "Functional and immunophenotypic modifications induced by interleukin-2 did not predict response to therapy in patients with renal cell carcinoma" <i>Journal of Biological Response Modifiers</i> 9:167-177 (1990).	

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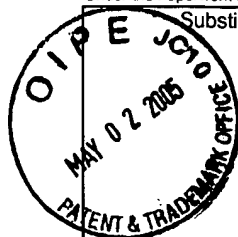
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		FERNANDES AND BALDWIN, "Interleukin-10 downregulates protective immunity to <i>Brucella abortus</i> " <i>Infect. Immun.</i> 63:1130-1133 (1995).	
		FOON, et al., "A prospective randomized trial of $\alpha_2\beta_2$ -interferon/ $\gamma$ -interferon or the combination in advanced metastatic renal cell carcinoma" <i>Journal of Biological Response Modifiers</i> 7:540-545 (1988).	
		FROST, et al., "Interleukin-6 induction by a muramyltripeptide derivative in cancer patients" <i>Journal of Biological Response Modifiers</i> 9:160-166 (1990).	
		GADDUCCI, et al., "Serum levels of soluble receptors for tumor necrosis factor (p55 and p75 sTNFr) in endometrial cancer" <i>Anticancer Res.</i> 16:3125-3128 (1996).	
		GILL, et al., "Interferon-alpha maintenance therapy after cytotoxic chemotherapy for treatment of acquired immunodeficiency syndrome-related kaposi's sarcoma" <i>Journal of Biological Response Modifiers</i> 9:512-516 (1990).	
		GREENBLATT, et al., "The type B receptor for tumor necrosis factor-alpha" <i>Blood</i> 80:1339-1346 (1992).	
		GUSTAVSON, et al., "Pharmacokinetics of teceleukin (Recombinant human inteleukin-2) after intravenous or subcutaneous administration to patients with cancer" <i>Journal of Biological Response Modifiers</i> 8:440-449 (1989).	
		HANDZEL, et al., "Immunomodulation of T cell deficiency in humans by thymic humoral factor: from crude extract to synthetic thymic humoral factor- $\gamma_2$ " <i>Journal of Biological Response Modifiers</i> 9:269-278 (1990).	
		HANK, et al., "Depressed <i>in vitro</i> T cell responses concomitant with augmented interleukin-2 responses by lymphocytes from cancer patients following <i>in vivo</i> treatment with interleukin-2" <i>Journal of Biological Response Modifiers</i> 9:5-14 (1990).	
		HERCEND, et al., "Immunotherapy with lymphokine-activated natural killer cells and recombinant interleukin-2: A feasibility trial in metastatic renal cell carcinoma" <i>Journal of Biological Response Modifiers</i> 9:546-555 (1990).	

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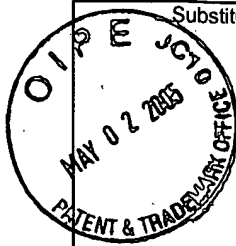
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		HERRMANN, et al., "Stimulation of granulopoiesis in patients with malignancy by recombinant human granulocyte-macrophage colony-stimulating factor: Assessment of two routes of administration" <i>Journal of Biological Response Modifiers</i> 9:475-479 (1990).	
		HERTLER, et al., "A phase I study of T101-ricin A chain immunotoxin in refractory chronic lymphocytic leukemia" <i>Journal of Biological Response Modifiers</i> 7:97-113 (1988).	
		HIMMLER, et al., "Molecular cloning and expression of human and rat tumor necrosis factor receptor chain (p60) and its soluble derivative, tumor necrosis factor-binding protein" <i>DNA and Cell Biol.</i> 9(10):705-715 (1990).	
		JACOBS, et al., "minimal antigenicity of intron A in human recipients demonstrated by three analytical methods" <i>Journal of Biological Response Modifiers</i> 7:447-456 (1988).	
		JAKOBSEN, et al., "Decreased antitoxic activities among children with clinical episodes of malaria" <i>Infect. Immun.</i> 66(4):1654-1659 (1998).	
		JAKSCHIES, et al., "Emergence and decay of the human Ms homolog in cancer patients during and after interferon- $\alpha$ therapy" <i>Journal of Biological Response Modifiers</i> 9:305-312 (1990).	
		KALMANTI, et al., "Serum levels of tumor necrosis factor soluble interleukin 2 receptor as markers of disease activity and prognosis in childhood leukemia and lymphoma" <i>Int. J. Hematol.</i> 57:147-152 (1993).	
		KAUFMANN, et al., "T cells and cytokines in intracellular bacterial infections: experiences with <i>Mycobacterium bovis</i> BCG" <i>Ciba Fdn. Symp.</i> 195:123-132 (1995).	
		KELLOKUMPU-LEHTINEN, et al., "Recombinant interferon- $\alpha$ 2a and vinblastine in advanced renal cell cancer: A clinical phase I-II study" <i>Journal of Biological Response Modifiers</i> 9:439-444 (1990).	
		KESSLER, "Adsorptive plasma treatment: Optimization of extracorporeal devices and systems" <i>Blood Purification</i> 11:150-157 (1993).	
		KHAZAELI, et al., "Initial evaluation of a human immunoglobulin M monoclonal antibody (HA-1A) in humans" <i>Journal of Biological Response Modifiers</i> 9:178-184 (1990).	

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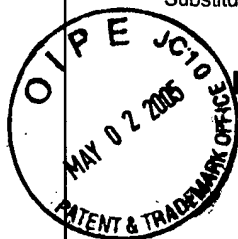
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		KOLITZ, et al., "Phase I trial of recombinant interleukin-2 and cyclophosphamide: Augmentation of cellular immunity and T-cell mitogenic response with long term administration of rIL-2" <i>Journal of Biological Response Modifiers</i> 7:457-472 (1988).	
		KRIGEL, et al., "Treatment of epidemic kaposi's sarcoma with a combination of interferon-alpha 2b and etoposide" <i>Journal of Biological Response Modifiers</i> 7:359-364 (1988).	
		LANTZ, et al., "Infusion of tumor necrosis factor (TNF) causes an increase in circulating TNF-binding protein in humans" <i>Cytokine</i> 2(6):402-406 (1990).	
		LASZLO, et al., "Phase I studies of recombinant interferon- $\gamma$ " <i>Journal of Biological Response Modifiers</i> 9:185-193 (1990).	
		LAUCELLA, et al., "Papel de las citoquinas en la resistencia y patologia durante la infeccion con <i>Trypanosoma cruzi</i> " <i>Revista Argentina de Microbiologia</i> , 28:99-109 (1996).	
		LENTZ, et al., "Apheresis of low molecular weight protein fraction and the onset of labor" <i>Journal of Clinical Apheresis</i> 5:62-67 (1990).	
		LENTZ, "The phylogeny of oncology" <i>Mol. Biother.</i> 2:137-144 (1990).	
		LETTERIO AND ROBERTS, "Regulation of immune responses by TGF- $\beta$ " <i>Ann. Rev. Immuno.</i> 16:137-161 (1998).	
		LITTON, et al., "Biological and clinical effects of the oral immunomodulator 3,6'Bis(2-piperidinoethoxy)acridine trihydrochloride in patients with malignancy" <i>Journal of Biological Response Modifiers</i> 9:61-70 (1990).	
		LUCEY, et al., "Type 1 and type 2 cytokine dysregulation in human infectious, neoplastic, and inflammatory diseases" <i>Clin. Micro. Rev.</i> 9(4):532-562 (1996).	
		MAAS, et al., "Interleukin-2 in cancer treatment: disappointing or (still) promising? A review" <i>Cancer Immunol. Immunother.</i> 36:141-148 (1993).	

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		MACA, "Inhibition of the growth of lewis lung carcinoma by indomethacin in conventional, nude, and beige mice" <i>Journal of Biological Response Modifiers</i> 9:568-580 (1990).	
		MARSHALL, et al., "Effects of coumarin (12-benzopyrone) and cimetidine on peripheral blood lymphocytes, natural killer cells, and monocytes in patients with advanced malignancies" <i>Journal of Biological Response Modifiers</i> 8:62-69 (1989).	
		MARSHALL, et al., "Treatment of renal cell carcinoma with daily low-dose alpha-interferon" <i>Journal of Biological Response Modifiers</i> 8:453-461 (1989).	
		MILES, et al., "induction of soluble tumour necrosis factor receptors during treatment with interleukin-2" <i>Brit. J. Cancer</i> 66:1195-1199 (1992).	
		MITTELMAN, et al., "Treatment of patients with advanced cancer using multiple long-term cultured lymphokine-activated killer (LAK) cell infusions and recombinant human interleukin-2" <i>Journal of Biological Response Modifiers</i> 8:468-478 (1989).	
		MUSIANI, et al., "Effect of low doses of interleukin-2 injected perilymphatically and peritumorally in patients with advanced primary head and neck squamous cell carcinoma" <i>Journal of Biological Response Modifiers</i> 8:571-578 (1989).	
		NEIDHART, "Phase I study of recombinant methionyl human consensus interferon (r-metHuIFN-Con)" <i>Journal of Biological Response Modifiers</i> 7:240-248 (1988).	
		ORATZ, et al. "Induction of tumor-infiltrating lymphocytes in human malignant melanoma metastases by immunization to melanoma antigen vaccine" <i>Journal of Biological Response Modifiers</i> 8:355-358 (1989).	
		ORATZ, et al., "Antimelanoma monoclonal antibody-ricin A chain immunoconjugate (XMMME-001-RTA) plus cyclophosphamide in the treatment of metastatic malignant melanoma: Results of a phase II trial" <i>Journal of Biological Response Modifiers</i> 9:345-354 (1990).	
		PAIS, et al., "Pharmacokinetics of recombinant interleukin-2 in children with malignancies: A pediatric oncology group study" <i>Journal of Biological Response Modifiers</i> 9:517-521 (1990).	

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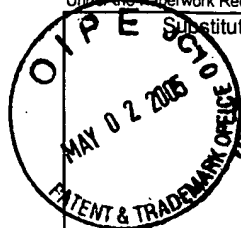
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		PAOLOZZI, et al., "Phase I trial of recombinant interleukin-2 and recombinant $\beta$ -interferon in refractory neoplastic diseases" <i>Journal of Biological Response Modifiers</i> 8:122-139 (1989).	
		PEREZ, et al., "A phase I trial of recombinant human gamma interferon (IFN- $\gamma_{4A}$ ) in patients with advanced malignancy" <i>Journal of Biological Response Modifiers</i> 7:309-317 (1988).	
		QUESADA, et al., "Recombinant interferon alpha and gamma in combination as treatment for tetastatic renal cell carcinoma" <i>Journal of Biological Response Modifiers</i> 7:234-239 (1988).	
		RABINOWITZ, et al., "Hemolytic anemia in a cancer patient treated with recombinant interferon- $\gamma$ " <i>Journal of Biological Response Modifiers</i> 9:256-259 (1990).	
		REIMANN, et al., "Suppression of the immune response by micro-organisms" <i>Scand. J. Immunol.</i> 31:543-546 (1990).	
		RIFFKIN, et al., "Defense against the immune barrage: helminth survival strategies" <i>Immunol. Cell Bio</i> 74:564-574 (1996)	
		ROMANI, et al., "T helper cell dichotomy to <i>Candida albicans</i> : implications for pathology" <i>Immunol. Res.</i> 14:148-162 (1995).	
		ROSENTHAL, et al., "The <i>in vitro</i> function of lymphocytes from 25 cancer patients receiving four to seven consecutive days of recombinant IL-2" <i>Journal of Biological Response Modifiers</i> 7:123-139 (1988).	
		RYBAK, et al., "Interferon therapy of relapsed and refractory hodgkin's disease: Cancer and leukemia group B study 8652" <i>Journal of Biological Response Modifiers</i> 9:1-4 (1990).	
		SARNA, et al., "Systemic administration of recombinant methionyl human interleukin-2 (Ala 125) to cancer patients: Clinical results" <i>Journal of Biological Response Modifiers</i> 8:16-24 (1989).	
		SARNA, et al., "A pilot study of intralymphatic interleukin-2. II. Clinical and biological effects" <i>Journal of Biological Response Modifiers</i> 9:81-86 (1990).	
		SARTHOU, et al., "Prognostic value of anti- <i>Plasmodium falciparum</i> -specific immunoglobulin G3, cytokines, and their soluble receptors in west African patients with severe malaria" <i>Infect. Immun.</i> 65(8):3271-3276 (1997).	

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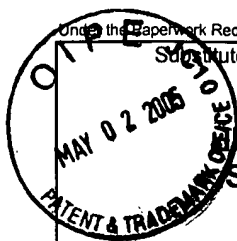
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Application Number	09/699,003
Filing Date	October 26, 2000
First Named Inventor	M. Rigdon Lentz
Group Art Unit	3762
Examiner Name	P. Bianco
Attorney Docket Number	LEN 101 CIP CON

Sheet	13	of	15
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		SATO, et al., "induction of bone formation in an adenoid cystic carcinoma of the maxillary sinus by adoptive immunotherapy involving intra-arterial injection of lympholine-activated killer cells and recombinant interleukin-2 in combination with radiotherapy" <i>Journal of Biological Response Modifiers</i> 9:329-334 (1990).	
		SCHAADT, et al., "Phase II study of recombinant human tumor necrosis factor in colorectal carcinoma" <i>Journal of Biological Response Modifiers</i> 9:247-250 (1990).	
		SCHALL, et al., "molecular cloning and expression of a receptor for human tumor necrosis factor" <i>Cell</i> 61:361-370 (1990).	
		SCHEITHAUER, et al., "Combined $\alpha$ -2C-interferon/VMCP polychemotherapy versus VMCP polychemotherapy a induction therapy in multiple myeloma: A prospective randomized trial" <i>Journal of Biological Response Modifiers</i> 8:109-115 (1989).	
		SCHILLER, et al., "A phase I trial of interferon- $\alpha$ -2a plus cyclophosphamide, vincristine, prednisone, and doxorubicin" <i>Journal of Biological Response Modifiers</i> 8:252-261 (1989).	
		SECKINGER, et al., "Purification and biologic characterization of a specific tumor necrosis factor $\alpha$ inhibitor" <i>J. Biol. Chem.</i> 264(20):11966-11973 (1989).	
		SEIGLER, et al, "Melanoma patient antibody responses to melanoma tumor-associated antigens defined by murine monoclonal antibodies" <i>Journal of Biological Response Modifiers</i> 8:37-52 (1989).	
		SHAU, et al., "A pilot study of intralymphatic interleukin-2. I. Cytotoxic and surface marker changes of peripheral blood lymphocytes" <i>Journal of Biological Response Modifiers</i> 9:71-80 (1990).	
		SIDHU AND BOLLON, "Tumor necrosis factor activities and cancer therapy – A perspective" <i>Pharmacol. Ther.</i> 57:79-128 (1993).	
		SELING, et al., "Immunosuppressive roles for IL-10 and IL-4 in human infection" <i>J. Immunol.</i> 150(12):5501-5510 (1993).	
		SPRIGGS, "One step ahead of the game: Viral immunomodulatory molecules" <i>Ann. Rev. Immunol.</i> 14:101-130 (1996).	

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		STEGER, et al., "Long-term remission in a patient with erythroleukemia following interferon- $\alpha$ treatment" <i>Journal of Biological Response Modifiers</i> 8:351-354 (1989).	
		STEINMETZ, et al., "Phase I study of 24-hour continuous intravenous infusion of recombinant human tumor necrosis factor" <i>Journal of Biological Response Modifiers</i> 7:417-423 (1988).	
		SZNOL, et al., "A phase I study of high-dose interleukin-2 in combination with interferon $\alpha_{2b}$ " <i>Journal of Biological Response Modifiers</i> 9:529-537 (1990).	
		TRIGG, et al., " $\alpha$ -interferon therapy for lymphoproliferative disorders developing in two children following bone marrow transplants" <i>Journal of Biological Response Modifiers</i> 8:603-613 (1989).	
		TRINCHIERI, et al., "Cytokine cross-talk between phagocytic cells and lymphocytes: Relevance for differentiation/Activation of phagocytic cells and regulation of adaptive immunity" <i>J. Cell. Biochem.</i> 53:301-308 (1993).	
		TRUMP et al., "Interferon- $\alpha$ -n1 and continuous infusion vinblastine for treatment of advanced renal cell carcinoma" <i>Journal of Biological Response Modifiers</i> 9:108-111 (1990).	
		UMIEL, et al., "Recombinant interleukin-2-activated intracavitary lymphocytes: Phenotypic characteristics and effector function" <i>Journal of Biological Response Modifiers</i> 8:409-421 (1989).	
		VON HOFF, et al., "Phase II evaluation of recombinant $\gamma$ -interferon in patients with advanced pancreatic carcinoma: A southwest oncology group study" <i>Journal of Biological Response Modifiers</i> 9:584-587 (1990).	
		WALSH, et al., "Phase I study of the combination of alpha-2 interferon and cisplatin" <i>Journal of Biological Response Modifiers</i> 8:11-15 (1989).	
		WEIL-HILLMAN, et al., "Transient decrease in IL-2 responsive lymphocytes 24 hours after initiation of continuous IL-2 infusion in cancer patients" <i>Journal of Biological Response Modifiers</i> 7:424-437 (1988).	

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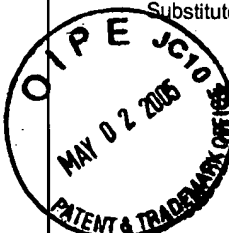
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		WHITEHEAD, et al., "A phase II trial of recombinant tumor necrosis factor in patients with metastatic colorectal adenocarcinoma: A southwest oncology group study" <i>Journal of Biological Response Modifiers</i> 9:588-591 (1990).	
		YELAVARTHI, et al., "Analysis of p60 and p80 tumor necrosis factor-alpha" <i>American Journal of Pathology</i> 143:1131-1141 (1993).	
		ZAMKOFF, et al., "A Phase I trial of subcutaneously administered recombinant tumor necrosis factor to patients with advanced malignancy" <i>Journal of Biological Response Modifiers</i> 8:539-552 (1989).	

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